## **Amendments to the Claims:**

This listing of claims will replace all prior versions, and listings, of claims in the application:

## **Listing of Claims:**

Claims 1-44 (Canceled).

45. (New) A method of cleaving a nucleic acid target in a cell comprising contacting the cell with a single-stranded siRNA molecule, wherein the single-stranded RNA molecule:

is complementary to the nucleic acid target molecule; is from 14 to 50 nucleotides in length; and

comprises a phosphate analog at the 5'-terminus; and

thereby cleaving the nucleic acid target molecule in the cell.

- 46. (New) The method of claim 45 wherein the single-stranded RNA molecule is from 15 to 29 nucleotides in length.
- 47. (New) The method of claim 45 wherein at least the 14 5'-terminal nucleotides of the single-stranded RNA molecule are complementary to the nucleic acid target molecule.

- 48. (New) The method of claim 45, wherein the phosphate analog at the 5'-terminus of the single-stranded RNA molecule is selected from among: a 5'-guanosine cap, a 5'-adenosine cap, a 5'-monothiophosphate, a 5'-monothiophosphate, a 5'-phosphorothiolate, a 5'-phosphoramidate, a 5'-alkyletherphosphonate.
- 49. (New) The method of claim 45, wherein the phosphate analog at the 5'-terminus of the single-stranded RNA molecule is selected from among: a 5'-monophosphate, a 5'-diphosphate, and a 5'-triphosphate.
- 50. (New) The method of claim 49, wherein the phosphate analog at the 5'-terminus of the single-stranded RNA molecule is a 5'-triphosphate.
- 51. (New) The method of claim 45, wherein the phosphate analog at the 5'-terminus of the single-stranded RNA molecule comprises a monophosphate, a diphosphate, or a triphosphate in which at least one oxygen atom of the monophosphate, diphosphate, or triphosphate has been replaced with a sulfur atom.
- 52. (New) The method of claim 51, wherein the phosphate analog is selected from among 5'-alpha-thiotriphosphate and 5'-gamma-thiotriphosphate.
- 53. (New) The method of claim 45, wherein the phosphate analog at the 5'-terminus of the single-stranded RNA molecule is an alkylphosphonate.

- 54. (New) The method of claim 53, wherein the alkylphosphonate has the formula: RP(OH)(O)-O-5' or  $(OH)_2(O)P-5$ '- $CH_2$ -, where R is a  $C_1$ - $C_3$  alkyl.
- 55. (New) The method of claim 45, wherein the phosphate analog at the 5'-terminus of the single-stranded RNA molecule is an alkyletherphosphonate.
- 56. (New) The method of claim 55, wherein the alkyletherphosphonate has the formula: RP(OH)(O)-O-5', where R is an alkylether.
- 57. (New) The method of claim 45, wherein the single-stranded RNA molecule comprises at least one modified nucleoside.
- 58. (New) The method of claim 57, wherein at least one modified nucleoside comprises a sugar modification.
- 59. (New) The method of claim 58, wherein at least one sugar modification is a 2'-sugar modification.
- 60. (New) The method of claim 45, wherein the single-stranded RNA molecule comprises at least one phosphorothioate linkage.

- 61. (New) The method of claim 45, wherein the single-stranded RNA molecule comprises at least one mismatch.
- 62. (New) The method of claim 45, wherein the single-stranded RNA molecule comprises a region at the 3'-terminus comprising at least one adenosine, guanosine or combination thereof.
  - 63. (New) The method of claim 45, wherein the cell is a eukaryotic cell.
- 64. (New) The method of claim 63, wherein the eukaryotic cell is a plant cell.
- 65. (New) The method of claim 63, wherein the eukaryotic cell is an animal cell.
- 66. (New) The method of claim 65, wherein the animal cell is selected from the group consisting of a mammalian cell, an embryonic cell, a pluripotent stem cell, a tumor cell and a virus-infected cell.
- 67. (New) The method of claim 66, wherein the tumor cell is a teratocarcinoma cell.

- 68. (New) The method of claim 65, wherein the animal cell is a human cell.
- 69. (New) A method of activating RISC and thereby cleaving a nucleic acid target molecule in a cell comprising contacting the cell with a single-stranded oligonucleotide, wherein the single-stranded oligonucleotide: is complementary to the nucleic acid target molecule; is from 15 to 29 nucleotides in length; and comprises a phosphate analog at the 5'-terminus; and thereby activating RISC and cleaving the nucleic acid target molecule in the cell.
- 70. (New) The method of claim 69 wherein at least the 14 5'-terminal nucleotides of the single-stranded oligonucleotide are complementary to the nucleic acid target molecule.
- 71. (New) The method of claim 69, wherein the phosphate analog at the 5'-terminus of the single-stranded oligonucleotide is selected from among: a 5'-guanosine cap, a 5'-adenosine cap, a 5'-monothiophosphate, a 5'-monodithiophosphate, a 5'-phosphorothiolate, a 5'-phosphoramidate, a 5'-alkyletherphosphonate.

- 72. (New) The method of claim 69, wherein the phosphate analog at the 5'-terminus of the single-stranded oligonucleotide is selected from among: a 5'-monophosphate, a 5'-diphosphate, and a 5'-triphosphate.
- 73. (New) The method of claim 72, wherein the phosphate analog at the 5'-terminus of the single-stranded oligonucleotide is a 5'-triphosphate.
- 74. (New) The method of claim 69, wherein the phosphate analog at the 5'-terminus of the single-stranded oligonucleotide comprises a monophosphate, a diphosphate, or a triphosphate in which at least one oxygen atom of the monophosphate, diphosphate, or triphosphate has been replaced with a sulfur atom.
- 75. (New) The method of claim 74, wherein the phosphate analog is selected from among 5'-alpha-thiotriphosphate and 5'-gamma-thiotriphosphate.
- 76. (New) The method of claim 69, wherein the phosphate analog at the 5'-terminus of the single-stranded RNA molecule is an alkylphosphonate.
- 77. (New) The method of claim 76, wherein the alkylphosphonate has the formula: RP(OH)(O)-O-5' or  $(OH)_2(O)P-5$ '- $CH_2$ -, where R is a  $C_1$ - $C_3$  alkyl.
- 78. (New) The method of claim 69, wherein the phosphate analog at the 5'-terminus of the single-stranded RNA molecule is an alkyletherphosphonate.

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- 79. (New) The method of claim 78, wherein the alkyletherphosphonate has the formula: RP(OH)(O)-O-5', where R is an alkylether.
- 80. (New) The method of claim 69, wherein the single-stranded oligonucleotide comprises at least one modified nucleoside.
- 81. (New) The method of claim 80, wherein at least one modified nucleoside comprises a sugar modification.
- 82. (New) The method of claim 81, wherein at least one sugar modification is a 2'-sugar modification.
- 83. (New) The method of claim 69, wherein the single-stranded RNA molecule comprises at least one phosphorothioate linkage.
- 84. (New) The method of claim 69, wherein the single-stranded RNA molecule comprises at least one mismatch.
- 85. (New) The method of claim 69, wherein the single-stranded RNA molecule comprises a region at the 3'-terminus comprising at least one adenosine, guanosine or combination thereof.

- 86. (New) The method of claim 69, wherein the cell is a eukaryotic cell.
- 87. (New) The method of claim 86, wherein the eukaryotic cell is a plant cell.
- 88. (New) The method of claim 86, wherein the eukaryotic cell is an animal cell.
- 89. (New) The method of claim 88, wherein the animal cell is selected from the group consisting of a mammalian cell, an embryonic cell, a pluripotent stem cell, a tumor cell and a virus-infected cell.
- 90. (New) The method of claim 89, wherein the tumor cell is a teratocarcinoma cell.
- 91. (New) The method of claim 89, wherein the animal cell is a human cell.
- 92. (New) The method of claim 45, wherein the single-stranded siRNA molecule is a single-stranded antisense siRNA molecule.